

SECTION V

REPORT ON CARCINOGENS

LISTING/DELISTING PROCEDURES

Nominations for listing or delisting an agent, substance, mixture or exposure circumstance in the Report on Carcinogens (RoC) should be submitted to the National Toxicology Program (NTP)¹. Nominations must contain a rationale for listing or delisting as either a “known human carcinogen” or a “reasonably anticipated human carcinogen”. Appropriate background information and relevant data (e.g. Journal articles, NTP Technical Reports, IARC listings, exposure surveys, release inventories, etc.) which support a nomination should be provided or referenced when possible.

An agent, substance, mixture or exposure circumstance nominated for listing or delisting will be evaluated initially by a National Institute of Environmental Health Sciences (NIEHS/NTP) RoC Review Committee (RG1), composed of scientists from the NIEHS/NTP, to determine if the information provided is sufficient to merit further consideration. If it is determined that the nomination contains insufficient information to warrant consideration by the NTP, it will be returned to the original nominator who will be invited to resubmit the nomination with additional justification, which may include new data, exposure information, etc. A notice, stating the action taken for a nominated substance found to contain insufficient justification for consideration, will be published in the Federal Register, trade journals, and NTP news updates, and included in subsequent editions of the RoC with the reason(s) it was not considered further. This decision will also be forwarded to the NTP Executive Committee² and the NTP Board of Scientific Counselors.

If it is determined the nomination warrants formal consideration it will be announced in the Federal Register, trade journals, and NTP publications to solicit public comment. The NTP may initiate an independent search of the literature and prepare a draft review document for the nomination under consideration. NTP draft documents will be prepared according to the following general format:

1.0 Introduction

1.1 Chemical Identification

synonyms, trade names, CAS #'s, molecular formula, molecular structure, etc.

1.2 Physical-Chemical Properties

1.3 Identification of Metabolites and Structural Analogs

2.0 Human Exposure

2.1 Use

2.2 Production

2.3 Analysis

2.4 Environmental Occurrence

environmental release, drinking water and food content, consumer products

2.5 Environmental fate

environmental fate in air, water, and soil

2.6 Environmental exposure

2.7 Occupational exposure

2.8 Biological indices of exposure

2.9 Regulations

Occupational Exposure Limits (standards and criteria), “other” standards and criteria

3.0 Human Studies

3.1 IARC studies

3.2 Case control studies

3.3 Cohort studies

4.0 Experimental Carcinogenesis

4.1 Previously reviewed studies

4.2 Recent Carcinogenicity Studies

5.0 Genotoxicity

6.0 Other Data Relevant to Evaluation of Carcinogenicity and its Mechanisms

Data used in the preparation of Sections 3 through 6 of the draft document must come from publicly available, peer reviewed sources.

FORMAL REVIEW STEPS

The following describes the review process for nominations that are considered by the NTP for listing in or delisting from the Report on Carcinogens:

NIEHS/NTP RoC Review Committee (RG1)

The original nomination and all public comments received in response to a nomination will be reviewed by RG1. Assignment of a primary and secondary reviewer will be made upon receipt of a nomination. Reviewers will lead discussions concerning the adequacy of the nomination. If the nomination warrants formal consideration, a search of pertinent databases will be performed and available citations will be reviewed by the primary reviewer. The primary reviewer will identify the relevant articles, and after consultation with the secondary reviewer, will obtain the identified literature. A draft summary of all available information from the original nomination and the literature search will be prepared. The primary and secondary reviewers will examine the nomination, the literature citations, and the draft document for completeness and adequacy. Public comments received in response to announcements of the nomination will be reviewed and issues identified will be addressed in the background document. The draft document will be revised if necessary and presented by the primary reviewer to the RG1. The nomination then continues through the review process.

Nominations reviewed by RG1 for which sufficient information could not be obtained will not proceed further. The subsequent RoC review groups, as well as the NTP Executive Committee, will be informed of this action and will have access to the information why it will not proceed further. The original nominator will be notified of the RG1 action and will be invited to resubmit the nomination with additional justification. All nominated agents, substances, or mixtures reviewed by RG1 but not selected for listing or delisting will be included in the subsequent edition of the RoC with the reason(s) why they were not considered further.

NTP Executive Committee's Interagency Working Group for the RoC (RG2)

The second review of nomination is by the NTP Executive Committee's Interagency Working Group. RG2 is a governmental interagency group that assesses whether relevant information on the nominated agent, substance, mixture or exposure circumstance is available and sufficient for listing in or delisting from the RoC. A reviewer for each nomination will be assigned from the RG2 and will be responsible for reviewing the draft document and for leading the RG2's discussion of the nomination. Public comments received in response to announcements of nominations will also be addressed by RG2 during the review. Upon completion of its review, RG2 will provide comments and recommendations for any changes and/or additions to the draft document and also make its recommendations for listing or delisting in the RoC.

Board of Scientific Counselors RoC Subcommittee (External Peer Review)

External peer review will be performed by a subcommittee of the NTP Board of Scientific Counselors. This subcommittee serves as an independent peer review group that assesses whether the relevant information available is sufficient for listing or delisting in the RoC. The subcommittee will review nominations in an open public meeting. Prior to public review, a notice will be published in the Federal Register, trade journals, and NTP publications, soliciting public comment. The notice will also invite interested groups or individuals to submit written comments and/or to address the subcommittee during the public review meeting. Reviewers for each nomination will be assigned from the subcommittee and will be responsible for reviewing the draft document and leading the subcommittee's discussion of the nomination. Upon completion of its review, the subcommittee will provide: comments and recommendations for any changes and/or additions to the draft document; and recommendations for listing or delisting the nominated agent, substance, mixture or exposure circumstance.

Public Comment

Upon completion of the reviews by RG1, RG2, and the Board subcommittee, those nominated agents, substances, or mixtures which are recommended for listing in or delisting from the RoC will be published in the Federal Register, trade journals, and NTP publications; and final public comment and input on the recommendations will be solicited.

NTP Executive Committee

The recommendations of RG1, RG2, and Board subcommittee, and all public comments received will be presented to the NTP Executive Committee² for review and comment.

NTP Director

The Director, NTP, reviews all recommendations and makes final decisions regarding the proposed listing and/or delisting and submits the RoC to the Office of the Secretary, DHHS. The Director, NTP, receives the three independent recommendations for the nominations from RG1, RG2, and the NTP Board RoC Subcommittee, and the consensus opinion of the NTP Executive Committee

concerning these recommendations. Based on this input and his evaluation of the relevant information, the Director makes his final recommendation of what nominations should be listed in or delisted from the latest edition of the draft RoC and submits it to the Secretary, HHS.

Secretary, Health and Human Service (HHS)

The final draft of the RoC is submitted by the Director, NTP to the Secretary, HHS for review and final approval. Upon approval of the RoC, the Secretary submits it to the U S Congress as a final document. The submission of the RoC to Congress constitutes publication of the Report and it becomes available to the public at that time. A notice of the publication and availability of the latest edition of the RoC, indicating all newly listed or delisted agents, substances, mixtures or exposure circumstances will be published in the Federal Register, trade journals and NTP newsletter publications.

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² Agencies represented on the NTP Executive Committee include:
Agency for Toxic Substances and Disease Registry (ATSDR), Consumer Product Safety Commission (CPSC),
Environmental Protection Agency (EPA), Food and Drug Administration (FDA), National Center for
Toxicological Research (NCTR), National Institute for Occupational Safety and Health (NIOSH), Occupational
Safety and Health Administration (OSHA), Department of Health and Human Services (DHHS), National
Institutes of Health (NIH), National Cancer Institute (NCI), and National Institute of Environmental Health
Sciences/NTP (NIEHS/NTP)